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37243C/21 SANKYO KK 03.10.78-JA-121801 (09.04.80) A61k-09/22 Preparing slow release pharmaceutical - by coating granules contg. active component with wax, water-sol. high mol. cpd. and nonionic surfactant and tableting	A96 B07 SANY 03.10.76 *J5 5049-312	A(12-VI) B(4-B1C, 4-C2, 4-C3, 12-M9, 12-M10, 12-M11). 4 etc. <u>EXAMPLE</u> None given.(7ppW5).
<p>Prepn. of slow release medicine comprises coating instantaneously disintegrating granules, which contain a major amt. of medicinal ingredient, with a film composed of wax, water-soluble high molecular compound and nonionic surfactant of HLB below 9, and tableting the coated granules.</p>		
<p><u>USE/ADVANTAGE</u> Granules which normally release the medicine in 1-10 minutes can be treated to release the medicine very slowly over 7-24 hours. Further the releasing velocity of the medicine is independent of pH and is almost constant.</p>		
<p><u>DETAILS</u> Usually a film of thickness 15-50 μ is coated on the granules. The water-soluble high molecular compound (the water solubility of which is independent of pH) can be methylcellulose, hydroxyethylcellulose, PVA, polyvinylpyrrolidone, hydroxypropylcellulose, polyethyleneglycol.</p>		

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EXHIBIT 30

PATENT ABSTRACTS OF JAPAN

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(54) PREPARATION OF SLOW-RELEASING MEDICINE

(57)Abstract:

PURPOSE: To obtain slow-releasing medicine which keeps the rate of release in the body constant irrespective of the pH in the digestive organs, by coating an easily disintegrable particle containing a principal medicine, with a layer which can easily be dissolved in body fluids, and forming the coated particles into a tablet.

CONSTITUTION: Easily disintegrable particle comprising a principal medicine and a disintegrating agent, is coated with a layer comprising wax, a water-soluble polymer and a nonionic surface-active agent having HLB of less than 9. The layer is water-soluble irrespective of the pH, and has a low water-permeability. The covered particles are formed into tablets having sufficiently high strength to endure the usual treatment. Although elution retarding rate of each covered particle is as short as 1W10min, a slow-releasing medicine which releases the effective component slowly, at a constant rate over a long period of 7W24hr can be made by forming into tablets.

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